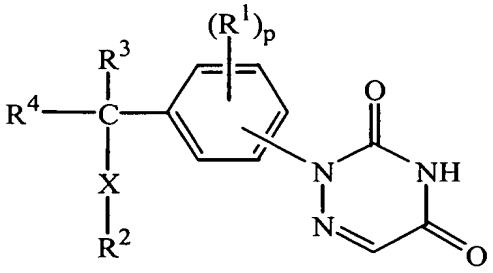
CLAIMS

1. A compound having the formula:



(I)

the N-oxides, the pharmaceutically acceptable addition salts and the stereochemically isomeric forms thereof, wherein :

p represents an integer being 0, 1, 2, 3 or 4;

X represents O, S, NR^5 or a direct bond or-X-R^2 taken together may represent cyano;

Y represents O, S, NR^5, or $S(O)_2$;

each R^1 independently represents C(=0)·Z-R^{14}, C_{1-6}alkyl, halo, polyhaloC_{1-6}alkyl, hydroxy, mercapto, C_{1-6}alkyloxy, C_{1-6}alkylthio, C_{1-6}alkylcarbonyloxy, aryl, cyano, nitro, Het^3, R^6, NR^7R^8 or C_{1-4}alkyl substituted with C(=0)-Z·R^{14}, Het^3, R^6 or NR^7R^8;

R^2 represents Het^1, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkyl or C_{1-6}alkyl substituted with one or two substituents selected from C(=0)·Z-R^{14}, hydroxy, mercapto, cyano, amino, mono- or di(C_{1-4}alkyl)amino, C_{1-6}alkyloxy optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkylthio optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkylsulfonyloxy, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, aryl, aryloxy, arylthio, Het^1, Het^1oxy and Het^1thio; and if X is O, S or NR^5, then R^2 may also represent aminothiocarbonyl, C_{1-4}alkylcarbonyl optionally substituted with C(=0)-Z-R^{14}, C_{1-4}alkylthiocarbonyl optionally substituted with C(=0)-Z-R^{14} , arylcarbonyl, arylthiocarbonyl, Het^1carbonyl or Het^1thiocarbonyl;

R^3 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl;

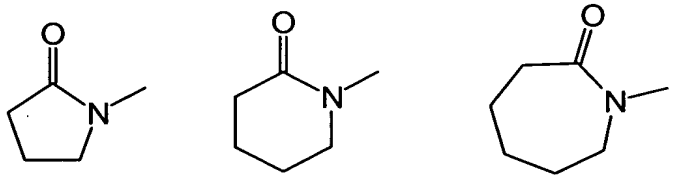
R^4 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl; or

R^3 and R^4 taken together form a C_{2-6}alkanediyl;

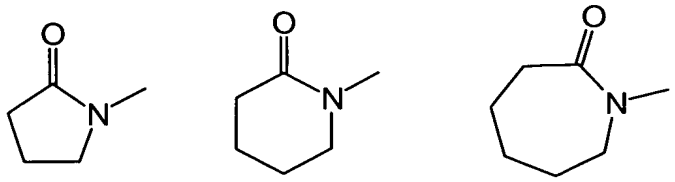
R^5 represents hydrogen or C_{1-4}alkyl;

each R^6 independently represents C_{1-6}alkylsulfonyl, aminosulfonyl, piperidinylsulfonyl, mono- or di(C_{1-4}alkyl)aminosulfonyl, mono- or di(benzyl)aminosulfonyl, polyhaloC_{1-6}alkylsulfonyl, C_{1-6}alkylsulfinyl, phenylC_{1-4}alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl, piperidinyl-aminosulfonyl, N-C_{1-4}alkyl-*N*-piperidinylaminosulfonyl, Y-R^{14}, mono- or di- (C_{1-4}alkyl)aminoC_{1-4}alkylsulfonyl, Het^6sulfonyl or C_{3-7} cycloalkylsulfonyl;

each R^7 and each R^8 are independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, mercapto-C_{1-4}alkyl, dihydroxyC_{1-4}alkyl, aryl, arylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, C_{1-4}alkyl-thiocarbonyl, arylcarbonyl, arylthiocarbonyl, Het^3thiocarbonyl, Het^3carbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6; or R^7 and R^8 taken together with the nitrogen atom to which they are attached form a radical of formula

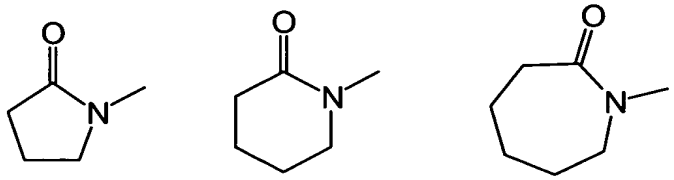


R^9 and R^{10} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, mercapto-C_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, arylcarbonyl, Het^3carbonyl, Het^3thiocarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6; or R^9 and R^{10} taken together with the nitrogen atom to which they are attached form a radical of formula



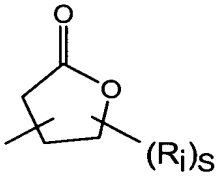
each R^{11} independently being selected from hydroxy, mercapto, cyano, nitro, halo, trihalomethyl, C_{1-4}alkyloxy optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkylthio optionally substituted with C(=0)-Z-R^{14}, formyl, trihaloC_{1-4}alkylsulfonyloxy, R^6, NR^7R^8, C(=O)$NR^{15}R^{16}$, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, aryl, aryloxy, arylcarbonyl, arylthiocarbonyl, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, C_{3-7}cycloalkyloxy optionally substituted with C(=0)-Z-R^{14}, C_{3-7}cycloalkylthio optionally substituted with C(=0)-Z-R^{14}, phthalimide-2-yl, Het^3 , Het^4, C(=O)Het^3, C(=O)C_{1-4}alkyl optionally be substituted with one or more substituents independently selected from hydroxy, mercapto, halo and phenyl;

R^{12} and R^{13} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, mercapto-C_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenyl-C_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, C_{1-4}alkylthiocarbonyl, arylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14} and R^6; or R^{12} and R^{13} taken together with the nitrogen atom to which they are attached form a radical of formula

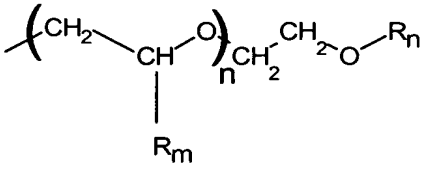


each R^{14} independently represents hydrogen; C_{1-20}acyl or C_{1-20}alkylC_{1-20}acyl (having a straight or branched, saturated or unsaturated hydrocarbon chain having 1 to 20 carbon atoms) optionally substituted with one or more substituents selected from hydroxy, mercapto, hydroxyC_{1-4}alkyl, mercapto-C_{1-4}alkyl, $NR^{17}R^{18}$, aryl, mono- or di-(C_{1-4}alkyl)amino, cyano and Het^5;

C_{1-20}alkyl optionally substituted with one or more substituents selected from hydroxy, halo, mercapto, C_{1-4}alkyloxyC_{1-4}alkyloxy, mercaptoC_{1-4}alkyl, $NR^{17}R^{18}$, aryl, mono- or di-(C_{1-4}alkyl)amino, cyano, Het^5, C_{1-4}alkyloxycarbonyl, arylC_{1-4} alkyloxycarbonyl, arylC_{1-4}alkyloxy, arylC_{1-4}alkylthiocarbonyl, arylC_{1-4}alkylthio, Het^5C_{1-4}alkyloxy, arylC_{1-4}alkylthio, C_{3-7} cycloalkyl and Het^5C_{1-4}alkylthio; C_{3-20}alkenyl optionally substituted with phenyl; C_{3-20}alkynyl; C_{3-7} cycloalkyl optionally substituted with one or more substituents selected from hydroxy, mercapto, halo, mercaptoC_{1-4}alkyl and hydroxyC_{1-4}alkyl; Het^5 or phenyl or R^{14} represents a radical having any of the following formulae:



(a)



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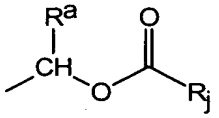

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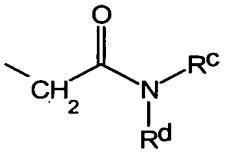
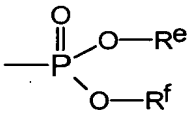


(d)



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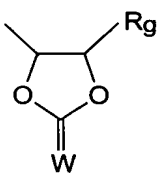

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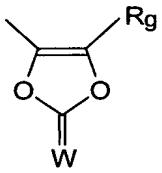

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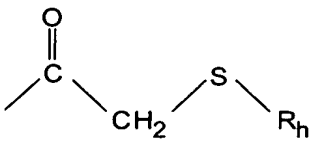

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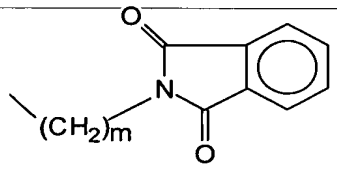

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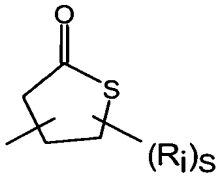

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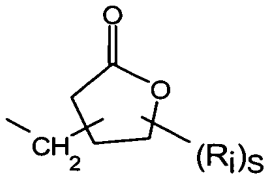

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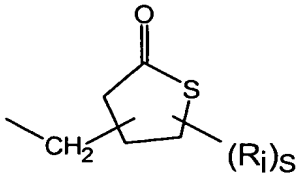
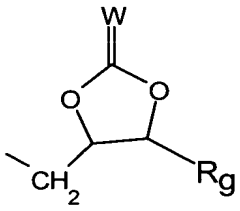
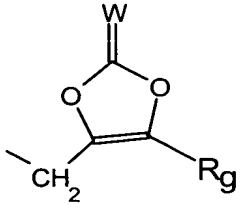


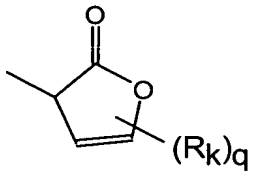
(n)



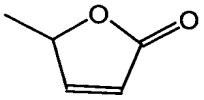
(o)



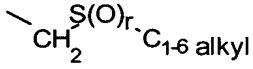
(p)



(q)



(r)



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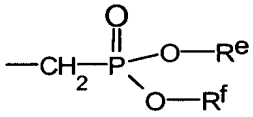


(t)

wherein m is 1 to 4, n is 0 to 5, q is 0 to 2, r is 0 to 2 and s is 0 to 4;

R^b is selected from hydrogen, C_{1-6}alkyl, phenyl, C_{3-7}cycloalkyl, C_{1-4} alkyloxyC_{1-6}alkyl and C_{1-4} alkyl-Y-C_{1-4}alkyl;

R^a, R^c, R^d, R^e and R^f are each independently selected from hydrogen, C_{1-6}alkyl, phenyl and C_{3-7}cycloalkyl, or R^e and R^f taken together may form $-CH_2-CH_2-$, $-CH_2-CH_2-CH_2-$ or $-CH_2-CH_2-CH_2-CH_2-$;

R_g, R_h and R_k are each independently hydrogen or C_{1-4} alkyl;

R_i is selected from hydroxy, C_{3-7}cycloalkyl and C_{1-4}alkyl, or two R_i taken together may form $-CH_2-CH_2-$, $-CH_2-CH_2-CH_2-$ or $-CH_2-CH_2-CH_2-CH_2-$ (thus building a spiro radical);

R_j is selected from $-O-R_b$; C_{1-6}alkyl optionally substituted with phenyl or

C_{3-7}cycloalkyl; phenyl; C_{3-7}cycloalkyl optionally substituted with C_{1-4} alkyloxy and mono-or di(C_{1-4}alkyl)amino;

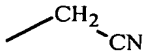
R_m is hydrogen or C_{1-4} alkyloxy;

R_n is hydrogen, C_{1-4}alkyl, C_{3-7}cycloalkyl, phenyl or phenylC_{1-4}alkyl; and

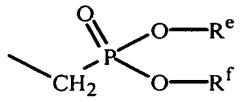
W represents O or S;

each Z independently represents O, S, NH, $-CH_2-O-$ or $-CH_2-S-$ whereby $-CH_2-$ is attached to the carbonyl group; or

$-Z-R^{14}$ taken together form a radical of formula

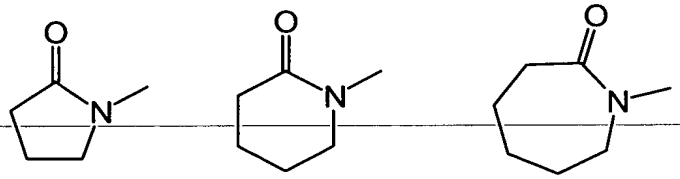


(f)



(g)

R^{15} and R^{16} are each independently selected from hydrogen; C_{1-4}alkyl optionally substituted with one or more substituents independently selected from hydroxy, mercapto, aryl, mono- or di(C_{1-4}alkyl) amino and pyridinyl; C_{1-4}alkyloxy; aryl; $-C(=O)-Z-R^{14}$; arylcarbonyl; arylthiocarbonyl; arylaminocarbonyl; arylaminothiocarbonyl; aminocarbonylmethylene; mono- or di(C_{1-4}alkyl) aminocarbonylmethylene; Het^{3}aminocarbonyl; Het^{3}aminothio-carbonyl; pyridinylC_{1-4}alkyl; Het^3 and R^6; or R^{15} and R^{16} taken together with the nitrogen atom to which they are attached form a radical of formula



R^{17} and R^{18} are each independently selected from hydrogen, C_{1-6}alkyl optionally substituted with one or more substituents independently selected from hydroxy, mercapto, aryl, mono- or di(C_{1-4}alkyl) amino, C_{1-4} alkyloxy and pyridinyl;

C_{1-4}alkyloxycarbonyl; aryl; C_{1-4}alkylcarbonyl; C_{1-4}alkylthiocarbonyl; arylcarbonyl; arylthiocarbonyl; arylaminocarbonyl; arylaminothiocarbonyl; C_{3-7}cycloalkyl;

C_{1-4}alkane-diyl-$C(=O)-Z-C_{1-6}$alkyl; $-C(=O)-Z-C_{1-6}$alkyl;

$-Y-C_{1-4}$alkanediyl-C(=O)-Z-C_{1-6}alkyl and R^6;

aryl represents phenyl optionally substituted with one, two or three substituents each independently selected from nitro, azido, cyano, halo, hydroxy, mercapto, C_{1-4}alkyl, C_{3-7}cycloalkyl, C_{1-4}alkyloxy, C_{1-4}alkylthio, formyl, polyhaloC_{1-4}alkyl, NR^9R^{10}, C(=O)NR^9R^{10}, C(=O)-Z-R^{14}, R^6, -O-R^6, phenyl, Het^3, C(=O)Het^3 and C_{1-4}alkyl substituted with one or more substituents each independently selected from halo, hydroxy, mercapto, C_{1-4}alkyloxy, C_{1-4}alkylthio, C(=O)-Z-R^{14}, $-Y-C_{1-4}$alkanediyl-C(=O)-Z-R^{14}, Het^3 or NR^9R^{10};

Het^1 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^2, R^{11} and C_{1-4}alkyl optionally substituted with one or, where possible, two or three substituents each independently selected from Het^2 and R^{11};

Het^2 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic

ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^4, R^{11} and C_{1-4}alkyl optionally substituted with one or, where possible, two or three substituents each independently selected from Het^4 and R^{11};

Het^3 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles include for instance pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxolanyl and tetrahydropyranyl; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C_{1-4}alkyl, C_{1-4}alkyloxy, C_{1-4}alkylcarbonyl, piperidinyl, $NR^{12}R^{13}$, $C(=O)$-Z-R^{14}, R^6 and C_{1-4}alkyl substituted with one or two substituents independently selected from hydroxy, carbonyl C_{1-4}alkyloxy, phenyl, $C(=O)$-Z-R^{14}, -Y-C_{1-4}alkanediyl-$C(=O)$-Z-R^{14}, R^6 and $NR^{12}R^{13}$;

Het^4 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles include for instance pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl,

thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and triazinyl;

Het^5 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic, monocyclic or polycyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus, or a fused polycyclic ring system including such heterocycle (such as for instance a fused benzoheterocycle); non-limiting examples of such heterocycles include for instance pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, tetrahydropyranyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, mercapto, carbonyl, C_{1-4}alkyl, C_{1-4}alkyloxy, C_{1-4}alkylthio, C_{1-4}alkylcarbonyl, piperidinyl, $NR^{17}R^{18}$, C(=O)-Z-C_{1-6}alkyl, R^6, sulfonamido and C_{1-4}alkyl substituted with one or two substituents independently selected from hydroxy, C_{1-4}alkyloxy, mercapto, C_{1-4}alkylthio, phenyl, C(=O)-Z-C_{1-6}alkyl, -Y-C_{1-4}alkanediyl-C(=O)-Z-C_{1-6}alkyl, R^6 and $NR^{17}R^{18}$;

Het^6 represents a three-membered, four-membered, five-membered or six-membered aromatic or non-aromatic monocyclic heterocycle comprising one or more, preferably one to four, heteroatoms, preferably selected from nitrogen, oxygen, sulfur and phosphorus; non-limiting examples of such heterocycles include for instance pyrrolidinyl, piperidinyl, azaridinyl, pyrazolinyl and pyrolinyl, wherein said heterocycle may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^2 , R^{11}

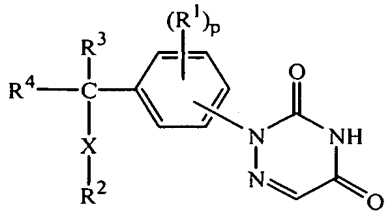
and C_{1-4}alkyl optionally substituted with one or more substituents independently selected from Het^2 and R^{11}.

provided however that

- R^2 is other than C_{1-6} alkyloxycarbonylC_{1-6}alkyl or aminocarbonyl; and
- R^7, R^8, R^9 and R^{10} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxyC_{1-4}alkylcarbonyl, C_{1-4}alkyloxycarbonylcarbonyl, C(=O)-O-R^{19}, C_{1-4}alkanediylC(=O)-O-R^{19} or –Y-C_{1-4}alkanediylC(=O)-O-R^{19}; and
- R^{12} and R^{13} are other than C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxyC_{1-4}alkylcarbonyl or C_{1-4}alkylcarbonylcarbonyl; and
- R^{11} is other than C(=O)-O-R^{19}, Y-C_{1-4}alkanediyl – C(=O)-OR^{19}, C(=O)NH_2, C(=O)NHC_{1-4}alkyl or C(=O)NHC_{3-7}cycloalkyl; and
- R^{15} and R^{16} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl or C_{1-4}alkyloxycarbonylcarbonyl; and
- aryl is other than phenyl substituted with C(=O)-O-R^{19}, C(=O)NH_2, C(=O)NHC_{1-4}alkyl, C(=O)NHC_{3-7}cycloalkyl and/or with C_{1-4}alkyl substituted with C(=O)-O-R^{19} or Y-C_{1-4}alkanediyl – C(=O)-O-R^{14}; and
- Het^3 is other than a monocyclic heterocycle substituted with C(=O)·O-R^{19} and/or with C_{1-4}alkyl substituted with C(=O)-O-R^{19} and/or Y-C_{1-4}alkanediyl C(=O)-O-R^{19}; and
- in each of the above proviso's R^{19} is defined as hydrogen, C_{1-4}alkyl, C_{3-7}cycloalkyl, aminocarbonylmethylene or mono- or di(C_{1-4}alkyl)aminocarbonylmethylene; and

wherein the said compound having the formula (I) contains at least one -C(=O)-Z-R^{14} moiety.

2. A compound according to claim 1 having the formula



(I')

a *N*-oxide, a pharmaceutically acceptable addition salt or a stereochemically isomeric form thereof, wherein :

p represents an integer being 0, 1, 2, 3 or 4;

X represents O, S, NR^5 or a direct bond or-X-R^2 taken together may represent cyano;

Y represents O, S, NR^5, or $S(O)_2$;

each R^1 independently represents C(=0)·Z-R^{14}, C_{1-6}alkyl, halo, polyhaloC_{1-6}alkyl, hydroxy, mercapto, C_{1-6}alkyloxy, C_{1-6}alkylthio, C_{1-6}alkylcarbonyloxy, aryl, cyano, nitro, Het^3, R^6, NR^7R^8 or C_{1-4}alkyl substituted with C(=0)-Z·R^{14}, Het^3, R^6 or NR^7R^8;

R^2 represents Het^1, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkyl or C_{1-6}alkyl substituted with one or two substituents selected from C(=0)·Z-R^{14}, hydroxy, cyano, amino, mono- or di(C_{1-4}alkyl)amino, C_{1-6}alkyloxy optionally substituted with C(=0)-Z-R^{14}, C_{1-6}alkylsulfonyloxy, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, aryl, aryloxy, arylthio, Het^1, Het^1oxy and Het^1thio; and if X is O, S or NR^5, then R^2 may also represent aminothiocarbonyl, C_{1-4}alkylcarbonyl optionally substituted with C(=0)-Z-R^{14}, C_{1-4}alkylthiocarbonyl optionally substituted with C(=0)-Z-R^{14} , arylcarbonyl, arylthiocarbonyl, Het^1carbonyl or Het^1thiocarbonyl;

R^3 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl;

R^4 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl; or

R^3 and R^4 taken together form a C_{2-6}alkanediyl;

R^5 represents hydrogen or C_{1-4}alkyl;

each R^6 independently represents C_{1-6}alkylsulfonyl, aminosulfonyl, mono- or di-(C_{1-4}alkyl)aminosulfonyl, mono- or di(benzyl)aminosulfonyl,

polyhaloC_{1-6}alkylsulfonyl, C_{1-6}alkylsulfinyl, phenylC_{1-4}alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl, piperidinylaminosulfonyl, *N*-C_{1-4}alkyl-*N*-piperidinylaminosulfonyl or mono-or di(C_{1-4}alkyl)aminoC_{1-4}alkylsulfonyl;

each R^7 and each R^8 are independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, aryl, arylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, arylcarbonyl, Het^3carbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6;

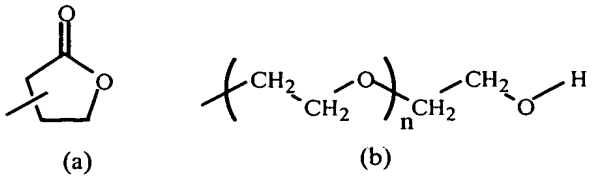
R^9 and R^{10} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, phenylcarbonyl, Het^3carbonyl, mono- or di(C_{1-4}alkyl)amino-C_{1-4}alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6;

each R^{11} independently being selected from hydroxy, mercapto, cyano, nitro, halo, trihalomethyl, C_{1-4}alkyloxy optionally substituted with C(=0)-Z-R^{14}, formyl, trihaloC_{1-4}alkylsulfonyloxy, R^6, NR^7R^8, C(=O)$NR^{15}R^{16}$, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, aryl, aryloxy, arylcarbonyl, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, C_{3-7}cycloalkyloxy optionally substituted with C(=0)-Z-R^{14}, phthalimide-2-yl, Het^3, Het^4 and C(=O)Het^3;

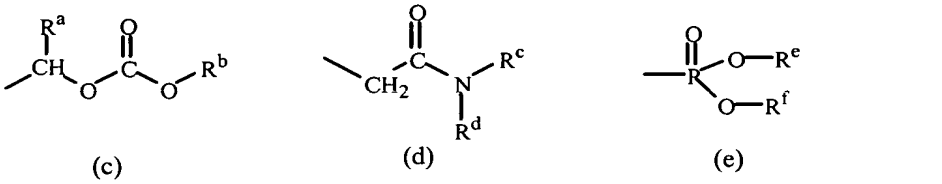
R^{12} and R^{13} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, phenylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14} and R^6;

each R^{14} independently represents C_{1-4} alkyl substituted with one or more substituents selected from phenyl, di- C_{1-4}alkylamino, cyano, Het^1 and C_{3-7} cycloalkyl, hydrogen, C_{1-20}acyl (having a straight or branched, saturated or

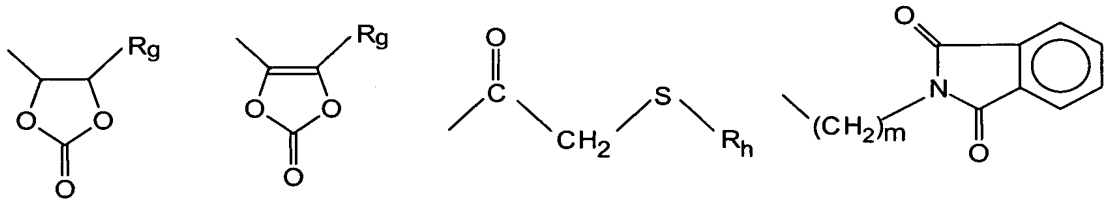
unsaturated hydrocarbon chain having 1 to 20 carbon atoms), C_{1-20}alkyl, C_{3-7}cycloalkyl, polyhaloC_{1-20}alkyl or a radical of formula



(a) (b)



(c) (d) (e)



(h) (i) (j) (k)

wherein n is 0 to 5 and m is 1 to 4;

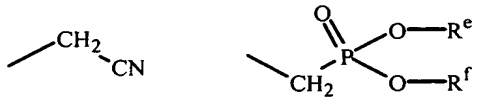
R^a, R^b, R^c, R^d, R^e and R^f are each independently hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl; or

R^e and R^f taken together may form $-CH_2-CH_2-$, $-CH_2-CH_2-CH_2-$ or $-CH_2-CH_2-CH_2-CH_2-$;

R_g and R_h are each independently C_{1-4} alkyl;

each Z independently represents O, S, NH, $-CH_2-O-$ or $-CH_2-S-$ whereby $-CH_2-$ is attached to the carbonyl group;

$-Z-R^{14}$ taken together form a radical of formula



(f) (g)

R^{15} and R^{16} are each independently selected from dihydroxyC_{1-4}alkyl, aryl, arylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, $-C(=O)-Z-R^{14}$, arylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, pyridinylC_{1-4}alkyl, Het^3, Het^4 or R^6;

aminocarbonylmethylene or mono-or di(C_{1-4}alkyl)aminocarbonylmethylene;

aryl represents phenyl optionally substituted with one, two or three substituents each independently selected from nitro, azido, cyano, halo, hydroxy, C_{1-4}alkyl, C_{3-7}cycloalkyl, C_{1-4}alkyloxy, formyl, polyhaloC_{1-4}alkyl, NR^9R^{10}, $C(=O)NR^9R^{10}$, $C(=O)$-Z-R^{14}, R^6, -O-R^6, phenyl, Het^3, $C(=O)Het^3$ and C_{1-4}alkyl substituted with one or more substituents each independently selected from halo, hydroxy, C_{1-4}alkyloxy, $C(=O)$-Z-R^{14}, -Y-C_{1-4}alkanediyl-$C(=O)$-Z-R^{14}, Het^3 or NR^9R^{10};

Het^1 represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^2, R^{11} and C_{1-4}alkyl optionally substituted with one or two substituents independently selected from Het^2 and R^{11};

Het^2 represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be

substituted with one, or where possible, two or three substituents each independently selected from Het^4, R^{11} and C_{1-4}alkyl optionally substituted with one or two substituents independently selected from Het^4 and R11;

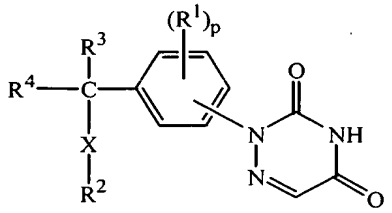
Het^3 represents a monocyclic heterocycle selected from pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl and tetrahydropyranyl; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C_{1-4}alkyl, C_{1-4}alkyloxy, C_{1-4}alkylcarbonyl, piperidinyl, $NR^{12}R^{13}$, C(=O)-Z-R^{14}, R^6 and C_{1-4}alkyl substituted with one or two substituents independently selected from hydroxy, C_{1-4}alkyloxy, phenyl, C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, R^6 and $NR^{12}R^{13}$;

Het^4 represents a monocyclic heterocycle selected from pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and triazinyl provided however that

- R^2 is other than C_{1-6} alkyloxycarbonylC_{1-6}alkyl, aminocarbonyl; and
- R^7, R^8, R^9 and R^{10} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy- C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl, C_{1-4}alkyloxycarbonylcarbonyl C(=O)-O-R^{14}, C_{1-4}alkanediylC(=O)-O-R^{14} and –Y-C_{1-4}alkanediylC(=O)-O-R^{14}; and
- R^{12} and R^{13} are other than C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl, C_{1-4}alkylcarbonylcabonyl; and
- R^{11} is other than C(=O)-O-R^{14}, Y-C_{1-4}alkanediyl – C(=O)-OR^{14}, $C(=O)NH_2$, $C(=O)NHC_{1-4}$alkyl or $C(=O)NHC_{3-7}$cycloalkyl; and
- R^{14} is other than hydrogen, C_{1-4}alkyl, C_{3-7}cycloalkyl, aminocarbonylmethylene, mono- or di (C_{1-4}alkyl) aminocarbonylmethylene in the event Z is 0; and
- R^{15} and R^{16} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl or C_{1-4}alkyloxycarbonylcarbonyl; and
- Aryl is other than phenyl substituted with C(=O)-O-R^{14} $C(=O)NH_2$, $C(=O)NHC_{1-4}$alkyl, $C(=O)NHC_{3-7}$cycloalkyl and/or with C_{1-4}alkyl substituted with C(=O)-O-R^{14} or Y-C_{1-4}alkanediyl – C(=O)-O-R^{14}; and

- Het^3 is other than a monocyclic heterocycle substituted with $C(=O)\cdot O\text{-}R^{14}$ and/or with C_{1-4}alkyl substituted with $C(=O)\text{-}O\text{-}R^{14}$ and/or $Y\text{-}C_{1-4}$alkanediyl – $(=O)\text{-}O\text{-}R^{14}$; and
- The said compound of formula (I) contains at least one – $C(=O)\text{-}Z\text{-}R^{14}$ moiety.

3. A compound according to claims 1 or 2 having the formula



(I")

a *N*-oxide, a pharmaceutically acceptable addition salt or a stereochemically isomeric form thereof, wherein :

p represents an integer being 0, 1, 2, 3 or 4;

X represents O, S, NR^5 or a direct bond or-$X\text{-}R^2$ taken together may represent cyano;

Y represents O, S, NR^5, or $S(O)_2$;

each R^1 independently represents $C(=0)\cdot Z\text{-}R^{14}$, C_{1-6}alkyl, halo, polyhaloC_{1-6}alkyl, hydroxy, mercapto, C_{1-6}alkyloxy, C_{1-6}alkylthio, C_{1-6}alkylcarbonyloxy, aryl, cyano, nitro, Het^3, R^6, NR^7R^8 or C_{1-4}alkyl substituted with $C(=0)\text{-}Z\cdot R^{14}$, Het^3, R^6 or NR^7R^8;

R^2 represents Het^1, C_{3-7}cycloalkyl optionally substituted with $C(=0)\text{-}Z\text{-}R^{14}$, C_{1-6}alkyl or C_{1-6}alkyl substituted with one or two substituents selected from $C(=0)\cdot Z\text{-}R^{14}$, hydroxy, cyano, amino, mono- or di(C_{1-4}alkyl)amino, C_{1-6}alkyloxy optionally substituted with $C(=0)\text{-}Z\text{-}R^{14}$, C_{1-6}alkylsulfonyloxy, C_{3-7}cycloalkyl optionally substituted with $C(=0)\text{-}Z\text{-}R^{14}$, aryl, aryloxy, arylthio, Het^1, Het^1oxy and Het^1thio; and if X is O, S or NR^5, then R^2 may also represent aminothiocarbonyl, C_{1-4}alkylcarbonyl optionally substituted with $C(=0)\text{-}Z\text{-}R^{14}$, C_{1-4}alkylthiocarbonyl optionally substituted with $C(=0)\text{-}Z\text{-}R^{14}$, arylcarbonyl, arylthiocarbonyl, Het^1carbonyl or Het^1thiocarbonyl;

R^3 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl;

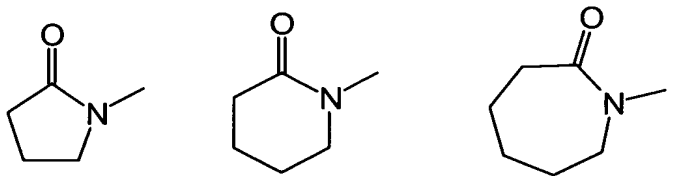
R^4 represents hydrogen, C_{1-6}alkyl or C_{3-7}cycloalkyl; or

R^3 and R^4 taken together form a C_{2-6}alkanediyl;

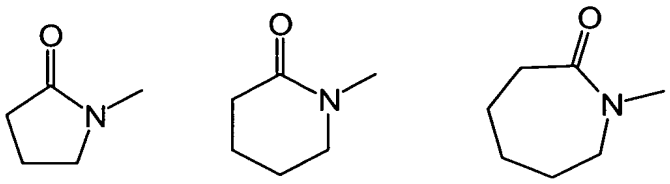
R^5represents hydrogen or C_{1-4}alkyl;

each R^6 independently represents C_{1-6}alkylsulfonyl, aminosulfonyl, piperidinylsulfonyl, mono- or di(C_{1-4}alkyl)aminosulfonyl, mono- or di(benzyl)aminosulfonyl, polyhaloC_{1-6}alkylsulfonyl, C_{1-6}alkylsulfinyl, phenylC_{1-4}alkylsulfonyl, piperazinylsulfonyl, aminopiperidinylsulfonyl, piperidinylaminosulfonyl, *N*-C_{1-4}alkyl-*N*-piperidinylaminosulfonyl or mono-or di(C_{1-4}alkyl)aminoC_{1-4}alkylsulfonyl;

each R^7 and each R^8 are independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, aryl, arylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, arylcarbonyl, Het^3carbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6; or R^7 and R^8 taken together with the nitrogen atom to which they are attached form a radical of formula

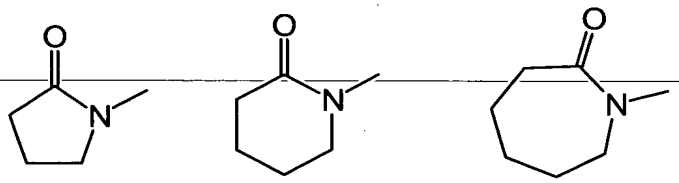


R^9 and R^{10} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, phenylcarbonyl, Het^3carbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, Het^3aminocarbonyl, Het^3aminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3, Het^4 and R^6; or R^9 and R^{10} taken together with the nitrogen atom to which they are attached form a radical of formula



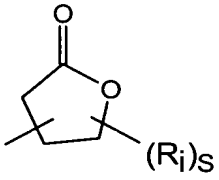
each R^{11} independently being selected from hydroxy, mercapto, cyano, nitro, halo, trihalomethyl, C_{1-4}alkyloxy optionally substituted with C(=0)-Z-R^{14}, formyl, trihaloC_{1-4}alkylsulfonyloxy, R^6, NR^7R^8, $C(=O)NR^{15}R^{16}$, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, aryl, aryloxy, arylcarbonyl, C_{3-7}cycloalkyl optionally substituted with C(=0)-Z-R^{14}, C_{3-7}cycloalkyloxy optionally substituted with C(=0)-Z-R^{14}, phthalimide-2-yl, Het^3, Het^4 and $C(=O)Het^3$;

R^{12} and R^{13} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, phenylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-R^{14}, -C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14} and R^6; or R^{12} and R^{13} taken together with the nitrogen atom to which they are attached form a radical of formula

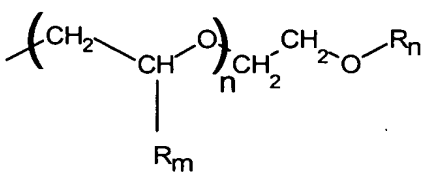


each R^{14} independently represents hydrogen, C_{1-20}acyl (having a straight or branched, saturated or unsaturated hydrocarbon chain having 1 to 20 carbon atoms), C_{1-20}alkyl, C_{3-20}alkenyl optionally substituted with phenyl, C_{3-20}alkynyl, C_{3-7} cycloalkyl, polyhaloC_{1-20}alkyl, Het^5 , phenyl or C_{1-20} alkyl substituted with one or more substituents selected from hydroxy, $NR^{17}R^{18}$, phenyl, mono- or di-(C_{1-4}alkyl)amino, cyano, Het^5 , C_{1-4} alkyloxycarbonyl,

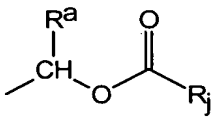
phenyl C_{1-4} alkyloxycarbonyl and C_{3-7} cycloalkyl, or R^{14} represents a radical of formula



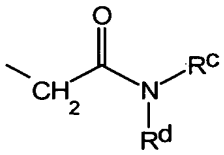
(a)



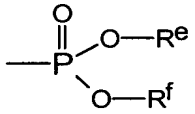
(b)



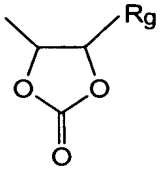
(c)



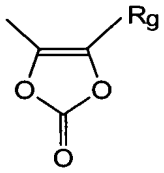
(d)



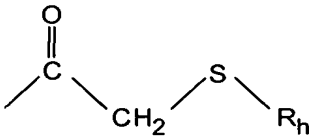
(e)



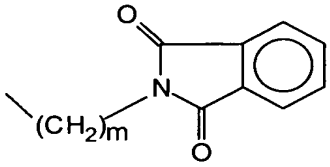
(h)



(i)

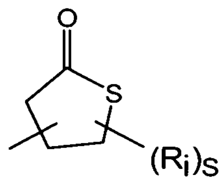


(j)

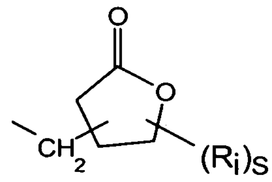


(k)

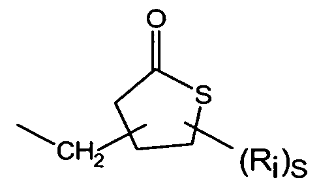
5



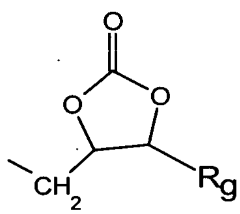
(l)



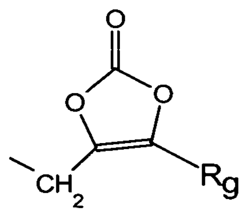
(m)



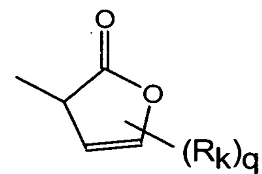
(n)



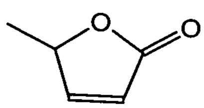
(o)



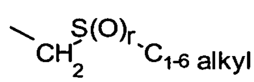
(p)



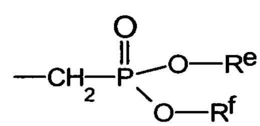
(q)



(r)



(s)



(t)

wherein m is 1 to 4, n is 0 to 5, q is 0 to 2, r is 0 to 2 and s is 0 to 4;

R^a, R^b, R^c, R^d, R^e and R^f are each independently hydrogen, $C_{1\text{-}6}$alkyl, phenyl or $C_{3\text{-}7}$cycloalkyl; or

R^e and R^f taken together may form $-CH_2\text{-}CH_2\text{-}$, $-CH_2\text{-}CH_2\text{-}CH_2\text{-}$ or $-CH_2\text{-}CH_2\text{-}CH_2\text{-}CH_2\text{-}$;

R_g, R_h and R_k are each independently hydrogen or $C_{1\text{-}4}$ alkyl;

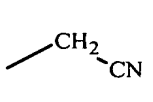
R_i is $C_{1\text{-}4}$alkyl;

R_j is $-O\text{-}R_b$, $C_{1\text{-}6}$alkyl, phenyl or $C_{3\text{-}7}$cycloalkyl optionally substituted with $C_{1\text{-}4}$ alkyloxy;

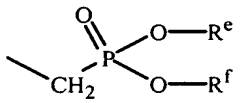
where R_m is hydrogen or $C_{1\text{-}4}$ alkyloxy and R_n is hydrogen, $C_{1\text{-}4}$alkyl, $C_{3\text{-}7}$cycloalkyl, phenyl or phenyl$C_{1\text{-}4}$alkyl

each Z independently represents O, S, NH, $-CH_2-O-$ or $-CH_2-S-$ whereby $-CH_2-$ is attached to the carbonyl group; or

$-Z-R^{14}$ taken together form a radical of formula

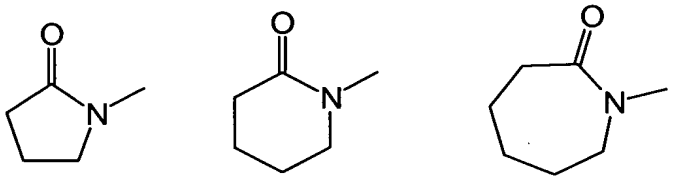


(f)



(g)

R^{15} and R^{16} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, aryl, arylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, $-C(=O)-Z-R^{14}$, arylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, arylaminocarbonyl, arylaminothiocarbonyl, aminocarbonylmethylene, mono- or di(C_{1-4}alkyl) aminocarbonylmethylene, Het^3aminocarbonyl, Het^3aminothiocarbonyl, pyridinylC_{1-4}alkyl, Het^3 or R^6; or R^{15} and R^{16} taken together with the nitrogen atom to which they are attached form a radical of formula



R^{17} and R^{18} are each independently selected from hydrogen, C_{1-4}alkyl, hydroxyC_{1-4}alkyl, dihydroxyC_{1-4}alkyl, phenyl, phenylC_{1-4}alkyl, C_{1-4}alkyloxyC_{1-4}alkyl, C_{1-4}alkylcarbonyl, phenylcarbonyl, mono- or di(C_{1-4}alkyl)aminoC_{1-4}alkyl, phenylaminocarbonyl, phenylaminothiocarbonyl, C_{3-7}cycloalkyl, pyridinylC_{1-4}alkyl, C_{1-4}alkanediyl-C(=O)-Z-C_{1-6}alkyl, -C(=O)-Z-C_{1-6}alkyl, -Y-C_{1-4}alkanediyl-C(=O)-Z-C_{1-6}alkyl and R^6;

aryl represents phenyl optionally substituted with one, two or three substituents each independently selected from nitro, azido, cyano, halo, hydroxy, C_{1-4}alkyl, C_{3-7}cycloalkyl, C_{1-4}alkyloxy, formyl, polyhaloC_{1-4}alkyl, NR^9R^{10}, $C(=O)NR^9R^{10}$, $C(=O)-Z-R^{14}$, R^6, $-O-R^6$, phenyl, Het^3, $C(=O)Het^3$ and C_{1-4}alkyl substituted with one or more substituents each independently selected from

halo, hydroxy, C_{1-4}alkyloxy, C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, Het^3 or NR^9R^{10};

Het^1 represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^2, R^{11} and C_{1-4}alkyl optionally substituted with one or two substituents independently selected from Het^2 and R^{11};

Het^2 represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^4, R^{11} and C_{1-4}alkyl optionally substituted with one or two substituents independently selected from Het^4 and R^{11};

Het^3 represents a monocyclic heterocycle selected from pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl and tetrahydropyranyl; wherein said monocyclic heterocycles each independently may optionally be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C_{1-4}alkyl, C_{1-4}alkyloxy, C_{1-4}alkylcarbonyl, piperidinyl, $NR^{12}R^{13}$, C(=O)-Z-R^{14}, R^6 and C_{1-4}alkyl substituted with one or two substituents independently selected from hydroxy, C_{1-4}alkyloxy, phenyl, C(=O)-Z-R^{14}, -Y-C_{1-4}alkanediyl-C(=O)-Z-R^{14}, R^6 and $NR^{12}R^{13}$;

Het^4 represents a monocyclic heterocycle selected from pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and triazinyl;

Het^5 represents a heterocycle selected from pyrrolyl, pyrrolinyl, imidazolyl, imidazolinyl, pyrazolyl, pyrazolinyl, triazolyl, tetrazolyl, furanyl, tetrahydrofuranyl, thienyl, thiolanyl, dioxolanyl, oxazolyl, oxazolinyl, isoxazolyl, thiazolyl, thiazolinyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, tetrahydropyranyl, dioxanyl, dithianyl, trithianyl, triazinyl, benzothienyl, isobenzothienyl, benzofuranyl, isobenzofuranyl, benzothiazolyl, benzoxazolyl, benzodioxanyl, indolyl, isoindolyl, indolinyl, purinyl, 1*H*-pyrazolo[3,4-d]pyrimidinyl, benzimidazolyl, quinolyl, isoquinolyl, cinnolinyl, phtalazinyl, quinazolinyl, quinoxalinyl, thiazolopyridinyl, oxazolopyridinyl and imidazo[2,1-b]thiazolyl; wherein said heterocycles each independently may be substituted with, where possible, one, two, three or four substituents each independently selected from hydroxy, C_{1-4}alkyl, C_{1-4}alkyloxy, C_{1-4}alkylcarbonyl, piperidinyl, $NR^{17}R^{18}$, C(=O)-Z-C_{1-6}alkyl, R^6, sulfonamido and C_{1-4}alkyl substituted with one or two substituents independently selected from hydroxy, C_{1-4}alkyloxy, phenyl, C(=O)-Z-C_{1-6}alkyl, -Y-C_{1-4}alkanediyl-C(=O)-Z-C_{1-6}alkyl, R^6 and $NR^{17}R^{18}$;

provided however that

- R^2 is other than C_{1-6} alkyloxycarbonylC_{1-6}alkyl or aminocarbonyl; and

- R^7, R^8, R^9 and R^{10} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl, C_{1-4}alkyloxycarbonylcarbonyl, C(=O)-O-R^{19}, C_{1-4}alkanediylC(=O)-O-R^{19} or –Y-C_{1-4}alkanediylC(=O)-O-R^{19}; and
- R^{12} and R^{13} are other than C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl or C_{1-4}alkylcarbonylcarbonyl; and
- R^{11} is other than C(=O)-O-R^{19}, Y-C_{1-4}alkanediyl – C(=O)-OR^{19}, C(=O)NH_2, C(=O)NHC_{1-4}alkyl or C(=O)NHC_{3-7}cycloalkyl; and
- R^{15} and R^{16} are other than aminocarbonyl, C_{1-4}alkylcarbonyloxy-C_{1-4}alkylcarbonyl, hydroxy C_{1-4}alkylcarbonyl or C_{1-4}alkyloxycarbonylcarbonyl; and
- aryl is other than phenyl substituted with C(=O)-O-R^{19}, C(=O)NH_2, C(=O)NHC_{1-4}alkyl, C(=O)NHC_{3-7}cycloalkyl and/or with C_{1-4}alkyl substituted with C(=O)-O-R^{19} or Y-C_{1-4}alkanediyl – C(=O)-O-R^{14}; and
- Het^3 is other than a monocyclic heterocycle substituted with C(=O)·O-R^{19} and/or with C_{1-4}alkyl substituted with C(=O)-O-R^{19} and/or Y-C_{1-4}alkanediyl – (=O)-O-R^{19}; and
- in each of the above proviso's R^{19} is defined as hydrogen, C_{1-4}alkyl, C_{3-7}cycloalkyl, aminocarbonylmethylene or mono- or di(C_{1-4}alkyl)aminocarbonylmethylene; and
- the said compound of formula (I) contains at least one – C(=O)-Z-R^{14} moiety.

4. A compound according to any of claims 1 to 3 wherein the 6-azauracil moiety is in the para position relative to the carbon atom bearing the -X-R^2, R^3 and R^4 substituents.

5. A compound according to any of claims 1 to 4 wherein R^2 is a monocyclic heterocycle selected from pyrrolyl, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, thiadiazolyl, oxadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyranyl, pyridazinyl and

triazinyl, wherein said monocyclic heterocycles each independently may optionally be substituted with one, or where possible, two or three substituents each independently selected from Het^2, R^{11} and C_{1-4}alkyl optionally substituted with Het^2 or R^{11}.

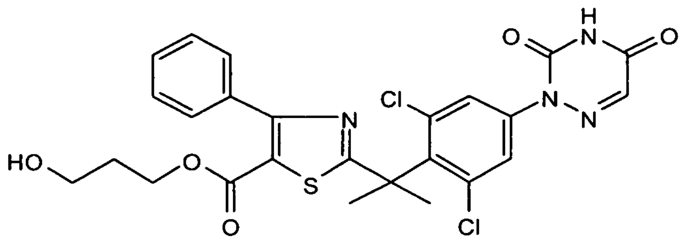
6. A compound according to any of claims 1 to 5 wherein R^3 and R^4 are both methyl and -X-R^2 is Het^1.

7. A compound according to any of claims 1 to 6 wherein p is 1 or 2 and each R^1 is chloro.

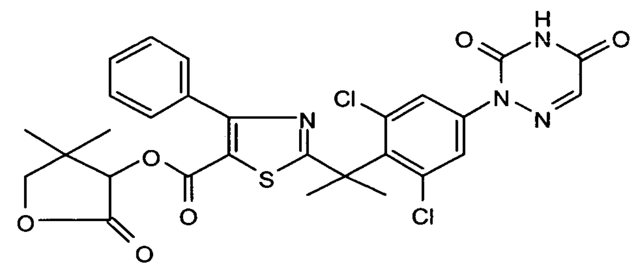
8 A compound according to any of claims 1 to 7 wherein R^3 and R^4 are both methyl, -X-R^2 is optionally substituted 2-thiazolyl or 3-oxadiazolyl, the 6-azauracil moiety is in the para position relative to the carbon atom bearing the -X-R^2, R^3 and R^4 substituents, and p is 2 whereby both R^1 substituents are chloro positioned ortho relative to the carbon atom bearing the -X-R^2, R^3 and R^4 substituents.

9 A compound according to claim 8 wherein X-R^2 is di-substituted with phenyl and either (i) R^{11} where R^{11} is a group of formula –C(=O)-Z-R^{14} in which Z is O and R^{14} is C_{1-20}alkyl substituted with hydroxy or with Het^5 where Het^5 is piperazinyl substituted with Het^6sulfonyl, or R^{14} is a radical of formula (a) in which R_j is C_{1-6}alkyl and s is 2, or (ii) C_{1-4}alkyl substituted with R^{11} where R^{11} is a a group of formula –C(=O)-Z-R^{14} in which Z is O and R^{14} is a radical of formula (a) in which R_j is C_{1-6}alkyl and s is 2.

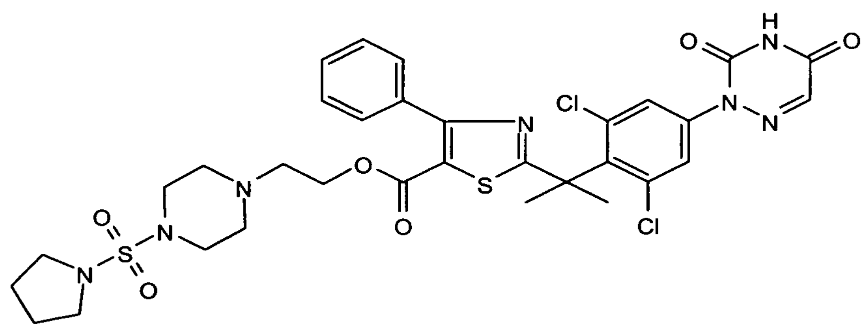
10 A compound according to claim 1 selected from those of formulae (A), (B), (C) and (D) below:-:



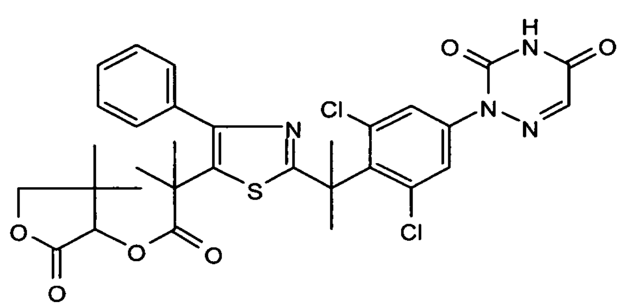
(A)



(B)



(C)



(D)

11. A composition comprising a pharmaceutically acceptable carrier and, as active ingredient, a therapeutically effective amount of a compound according to any of claims 1 to 10.

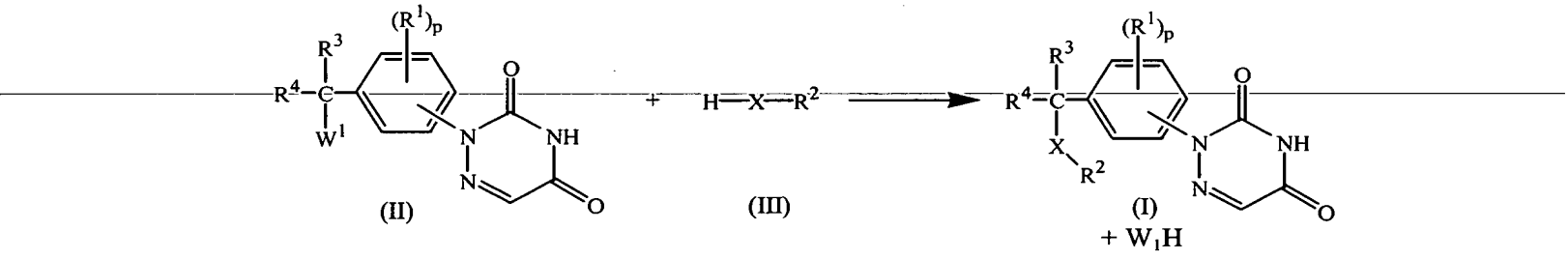
12. A process for preparing a composition as claimed in claim 11, wherein a pharmaceutically acceptable carrier is intimately mixed with a therapeutically effective amount of a compound according to any of claims 1 to 10.

13. A compound as claimed in any one of claims 1 to 10 for use as a medicine.

14. Use of a compound according to any of claims 1 to 10 in the manufacture of a medicament for treating eosinophil-dependent inflammatory diseases.

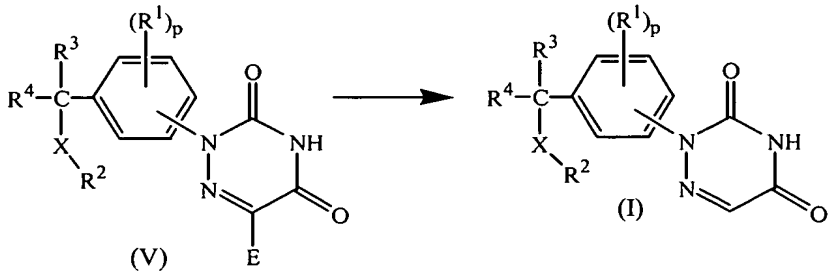
15. A process for preparing a compound as claimed in claim 1, comprising the step of

a) reacting an intermediate of formula (II) wherein W^1 is a suitable leaving group with an appropriate reagent of formula (III) optionally in a reaction-inert solvent and optionally in the presence of a base at a temperature ranging between –70°C and reflux temperature;



wherein R^1 R^2, R^3, R^4, p and X are as defined in claim 1 or;

b) eliminating the group E of a triazinedione of formula (V)



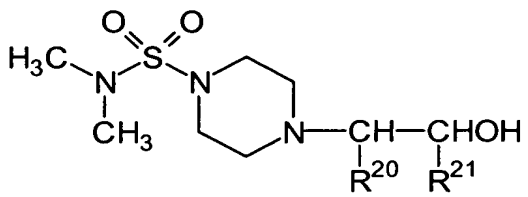
(V) (I)

wherein E is an appropriate electron attracting group and R^1, R^2, R^3, R^4, X and p are as defined in claim 1; and, if desired, converting compounds of formula (I) into each other following art-known transformations, and further, if desired, converting the compounds of formula (I), into a therapeutically active non-toxic acid addition salt by treatment with an acid, or into a therapeutically active non-toxic base addition salt by treatment with a base, or conversely, converting the acid addition salt form into the free base by treatment with alkali, or converting the base addition salt into the free acid by treatment with acid; and also, if desired, preparing stereochemically isomeric forms or *N*-oxide forms thereof.

16. A process of marking a receptor comprising the steps of
    a) radiolabelling a compound as defined in claim 1;
    b) administering said radiolabelled compound to biological material,
    c) detecting the emissions from the radiolabelled compound.

17 A process of imaging an organ, characterized by, administering a sufficient amount of a radiolabelled compound of formula (I) in an appropriate composition, and detecting the emissions from the radioactive compound.

18. A compound of formula

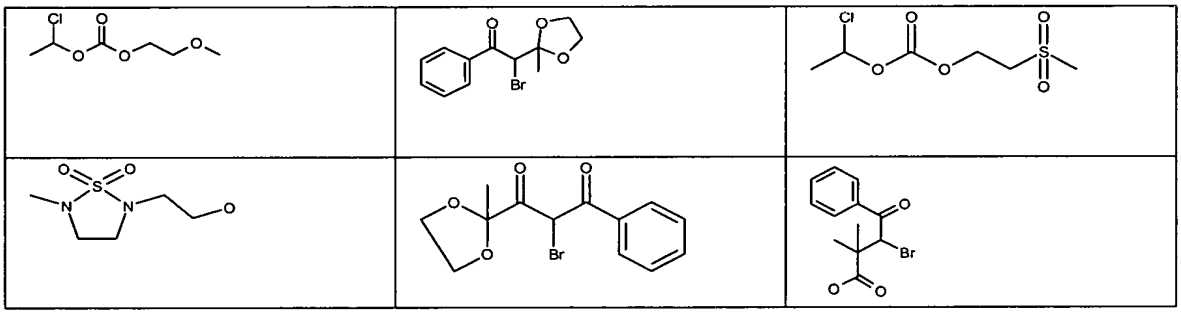


(XXIII)

wherein R^{20} and R^{21} are each independently selected from hydrogen or C_{1-20} alkyl or R^{20} and R^{21} taken together with the carbon atom to which they are attached form a cycloalkyl radical.

19. Use of a compound of claim 18 for preparing a compound of claim 1 wherein Het^5 represents a sulfonamido substituted piperazine.

20. A compound having any of the following formulae:



21. Use of a compound of claim 20 as an intermediate for preparing a compound of claim 1.